WEST Search History

Hide Items Restore Clear Cancel

DATE: Thursday, November 15, 2007

Hide? Set Name Query

Hit Count

DB=USPT; PLUR=YES; OP=ADJ

L1 (546/219.ccls. or 514/328.ccls.) and (depression or anxiety or apnoea or migraine)

40

END OF SEARCH HISTORY

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=> .s C19 H28 N2 O3/mf
          1376 C19 H28 N2 O3/MF
=> s 14 and piperidin?
       1236574 PIPERIDIN?
           277 L4 AND PIPERIDIN?
1.5
=> s 15 and piperidinedione
          4873 PIPERIDINEDIONE
             4 L5 AND PIPERIDINEDIONE
L6
=> d 1-3
     ANSWER 1 OF 4 REGISTRY COPYRIGHT 2007 ACS on STN
L6
RN
     732209-36-8 REGISTRY
ED
     Entered STN: 24 Aug 2004
     2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-
CN
     4,4-dimethyl-, (+)- (CA INDEX NAME)
FS
     STEREOSEARCH
MF
     C19 H28 N2 O3
CI
     COM
SR
     CA
Rotation (+).
          Мe
            Me
                     NMe<sub>2</sub>
  HN
             (CH<sub>2</sub>) 3
      റ
                OMe
**PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT**
L6
     ANSWER 2 OF 4 REGISTRY COPYRIGHT 2007 ACS on STN
RN
     117576-37-1 REGISTRY
ED
     Entered STN: 18 Nov 1988
CN
     2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-
     4,4-dimethyl-, (-)- (CA INDEX NAME)
FS
     STEREOSEARCH
MF
     C19 H28 N2 O3
CI
     COM
SR
     CA
LC
     STN Files:
                  BEILSTEIN*, CA, CAPLUS, USPATFULL
         (*File contains numerically searchable property data)
Rotation (-).
```

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

6 REFERENCES IN FILE CA (1907 TO DATE)

6 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L6 ANSWER 3 OF 4 REGISTRY COPYRIGHT 2007 ACS on STN

RN 117539-18-1 REGISTRY

ED Entered STN: 11 Nov 1988

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,5-dimethyl- (9CI) (CA INDEX NAME)

MF C19 H28 N2 O3

SR CA

LC STN Files: CA, CAPLUS

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1907 TO DATE)

1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

=> FILE REG

COST IN U.S. DOLLARS

SINCE FILE TOTAL

ENTRY SESSION

32.10 40.34

FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 11:02:53 ON 15 NOV 2007 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

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Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 14 NOV 2007 HIGHEST RN 953817-57-7 DICTIONARY FILE UPDATES: 14 NOV 2007 HIGHEST RN 953817-57-7

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH June 29, 2007

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REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

http://www.cas.org/support/stngen/stndoc/properties.html

=> STR 732209-36-8

:END

L7 STRUCTURE CREATED

=> S L7 FAM FUL

FULL SEARCH INITIATED 11:02:56 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 20 TO ITERATE

100.0% PROCESSED 20 ITERATIONS 13 ANSWERS

SEARCH TIME: 00.00.01

L8 13 SEA FAM FUL L7

=>

=> D SCAN

L8 13 ANSWERS REGISTRY COPYRIGHT 2007 ACS on STN

IN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4dimethyl-, sulfate (1:1)

MF C19 H28 N2 O3 . H2 O4 S

CM 1

CM 2

HOW MANY MORE ANSWERS DO YOU WISH TO SCAN? (1):12

L8 13 ANSWERS REGISTRY COPYRIGHT 2007 ACS on STN

IN 1H-1,4-Benzodiazepine-3-carboxylic acid, 7-chloro-2,3-dihydro-2-oxo-5phenyl-, monopotassium salt, compd. with potassium hydroxide (K(OH))
 (1:1), mixt. with 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4dimethyl-2,6-piperidinedione (9CI)

MF C19 H28 N2 O3 . C16 H11 Cl N2 O3 . H K O . K

CI MXS

CM 1

CM 2

CM 3

K

CM 4

к-он

L8 13 ANSWERS REGISTRY COPYRIGHT 2007 ACS on STN
IN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4dimethyl-, (+)MF C19 H28 N2 O3

CI COM

Rotation (+).

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

L8 13 ANSWERS REGISTRY COPYRIGHT 2007 ACS on STN

IN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4dimethyl-, (-)-

MF C19 H28 N2 O3

CI COM

Rotation (-).

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

L8 13 ANSWERS REGISTRY COPYRIGHT 2007 ACS on STN

IN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl-, monohydrochloride, (+)- (9CI)

MF C19 H28 N2 O3 . Cl H

Rotation (+).

● HCl

L8 13 ANSWERS REGISTRY COPYRIGHT 2007 ACS on STN

IN 2-Naphthalenecarboxylic acid, 4,4'-methylenebis[3-hydroxy-, compd. with 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl-2,6-piperidinedione (1:1) (9CI)

MF C23 H16 O6 . C19 H28 N2 O3

CM 1

CM 2

L8 13 ANSWERS REGISTRY COPYRIGHT 2007 ACS on STN

IN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4dimethyl-, monohydrochloride, mixt. with 7-chloro-1,3-dihydro-1-methyl-5phenyl-2H-1,4-benzodiazepin-2-one (9CI)

MF C19 H28 N2 O3 . C16 H13 Cl N2 O . Cl H

CI MXS

● HCl

CM 2

L8 13 ANSWERS REGISTRY COPYRIGHT 2007 ACS on STN

IN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4dimethyl-

MF C19 H28 N2 O3

CI COM

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

L8 13 ANSWERS REGISTRY COPYRIGHT 2007 ACS on STN

IN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4dimethyl-, hydrochloride (9CI)

MF C19 H28 N2 O3 . \times Cl H

Me (CH₂)
$$_3$$
 – NMe₂

•x HCl

L8 13 ANSWERS REGISTRY COPYRIGHT 2007 ACS on STN

IN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4dimethyl-, monohydrochloride, (-)- (9CI)

MF C19 H28 N2 O3 . Cl H

Rotation (-).

● HCl

CM 1

Rotation (-).

CM 2

L8 13 ANSWERS REGISTRY COPYRIGHT 2007 ACS on STN

IN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl-, monohydrochloride, mixt. with 7-chloro-N-methyl-5-phenyl-3H-1,4-benzodiazepin-2-amine 4-oxide, monohydrochloride (9CI)

MF C19 H28 N2 O3 . C16 H14 Cl N3 O . 2 Cl H

CI MXS

CM 1

HCl

CM 2

HC1

L8 13 ANSWERS REGISTRY COPYRIGHT 2007 ACS on STN

IN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4dimethyl-, hydrochloride (1:1)

MF C19 H28 N2 O3 . Cl H

CI COM

HCl

ALL ANSWERS HAVE BEEN SCANNED

=> FILE REG

COST IN U.S. DOLLARS

SINCE FILE TOTAL

ENTRY SESSION

68.15 108.49

FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 11:04:17 ON 15 NOV 2007 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2007 American Chemical Society (ACS)

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New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH June 29, 2007

Please note that search-term pricing does apply when conducting SmartSELECT searches.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

http://www.cas.org/support/stngen/stndoc/properties.html

=> STR 117576-37-1

WARNING. STEREO DATA NOT INCLUDED IN MODEL (NOT SEARCHABLE): END

L9 STRUCTURE CREATED

=> S L9 FAM FUL

FULL SEARCH INITIATED 11:04:21 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 20 TO ITERATE

100.0% PROCESSED 20 ITERATIONS 13 ANSWERS

SEARCH TIME: 00.00.01

L10 13 SEA FAM FUL L9

(FILE 'HOME' ENTERED AT 10:55:56 ON 15 NOV 2007)

FILE 'REGISTRY' ENTERED AT 10:57:19 ON 15 NOV 2007 L1 1 S AGN 2979/CN

FILE 'CAPLUS' ENTERED AT 10:57:57 ON 15 NOV 2007

FILE 'REGISTRY' ENTERED AT 10:58:12 ON 15 NOV 2007

L2 1 S 53873-28-2 L3 1 S 117539-17-0

L4 1376 S C19 H28 N2 O3/MF L5 277 S L4 AND PIPERIDIN?

L5 277 S L4 AND PIPERIDIN? L6 4 S L5 AND PIPERIDINEDIONE

FILE 'REGISTRY' ENTERED AT 11:02:53 ON 15 NOV 2007

L7 STR 732209-36-8

L8 13 S L7 FAM FUL

FILE 'REGISTRY' ENTERED AT 11:04:17 ON 15 NOV 2007

L9 STR 117576-37-1

L10 13 S L9 FAM FUL

=> fil caplus

COST IN U.S. DOLLARS

SINCE FILE TOTAL

ENTRY SESSION 68.15 176.64

FULL ESTIMATED COST

FILE 'CAPLUS' ENTERED AT 11:05:30 ON 15 NOV 2007 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2007 AMERICAN CHEMICAL SOCIETY (ACS)

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FILE COVERS 1907 - 15 Nov 2007 VOL 147 ISS 21 FILE LAST UPDATED: 14 Nov 2007 (20071114/ED)

Effective October 17, 2005, revised CAS Information Use Policies apply. They are available for your review at:

http://www.cas.org/infopolicy.html

=> s 110

L11 19 L10

=> s 11

L12 15 L1

=> s 111 not 112

L13 4 L11 NOT L12

=> d bib abs hitstr 1-4

L13 ANSWER 1 OF 4 CAPLUS COPYRIGHT 2007 ACS on STN

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2004:182707 CAPLUS
ΝA
DΝ
     140:210809
     Piperidin-2,6-dione pamoate salts for the treatment of stress-related
TI
     affective disorders, and pharmaceutical compositions containing them
IN
     Wermuth, Camille Georges
PA
     Prestwick Pharmaceuticals, Inc., USA
     PCT Int. Appl., 21 pp.
SO
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 1
     PATENT NO.
                         KIND
                                DATE
                                            APPLICATION NO.
                                                                    DATE
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     WO 2004017970
                                20040304
                                            WO 2003-IB3698
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PRAI GB 2002-19639
                          Α
                                20020822
     WO 2003-IB3698
                                20030818
                          W
     MARPAT 140:210809
OS
```

$$R^{1}$$
 R^{2}
 $(CH_{3})_{2}N-(CH_{2})_{n}$
 N

GI

AB Pamoate salts of certain 3-phenyl-3-dimethylaminoalkyl-4,4dimethylpiperidin-2,6-diones,(I) (R1 = MeO, EthO, OH; R2 = MeO, EthO, OH; n = 2, 3) and pharmacol. acceptable solvates thereof are devoid of the weight loss and hepatocyte changes in the rat which limited to marginally effective levels the permitted clin. doses of the corresponding

I

hydrochlorides in the treatment or prophylaxis of stress-related affective disorders such as anxiety, depression, migraine and sleep apnea. The preferred pamoate salts are 3(3,5dimethoxyphenyl)-3-(3-dimethylaminopropyl)-4,4-dimethylpiperidine-2,6-dione pamoate and, especially, 3(3-methoxyphenyl)-3-(3-dimethylaminopropyl)-4,4-dimethylpiperidine2,6-dione pamoate.

IT 666175-71-9P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(piperidin-2,6-dione pamoate salts for treatment of stress-related affective disorders, and pharmaceutical compns. containing them)

RN 666175-71-9 CAPLUS

2-Naphthalenecarboxylic acid, 4,4'-methylenebis[3-hydroxy-, compd. with 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl-2,6-piperidinedione (1:1) (9CI) (CA INDEX NAME)

CM 1

CN

CRN 53873-21-5 CMF C19 H28 N2 O3

CM 2

CRN 130-85-8 CMF C23 H16 O6

IT 666175-73-1

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(piperidin-2,6-dione pamoate salts for treatment of stress-related affective disorders, and pharmaceutical compns. containing them)

RN 666175-73-1 CAPLUS

CN 2-Naphthalenecarboxylic acid, 4,4'-methylenebis[3-hydroxy-, compd. with (-)-3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl-2,6-piperidinedione (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 117576-37-1 CMF C19 H28 N2 O3

Rotation (-).

CM 2

CRN 130-85-8 CMF C23 H16 O6

IT 500350-77-6P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(piperidin-2,6-dione pamoate salts for treatment of stress-related affective disorders, and pharmaceutical compns. containing them)

RN 500350-77-6 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl-, sulfate (1:1) (CA INDEX NAME)

CM 1

CRN 53873-21-5 CMF C19 H28 N2 O3

Me Me OMe
$$(CH_2)_3 - NMe_2$$

CM2

CRN 7664-93-9 CMF H2 O4 S

RE.CNT 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 2 OF 4 CAPLUS COPYRIGHT 2007 ACS on STN

AN 2003:174547 CAPLUS

DN 138:204953

Preparation of piperidine-2,6-dione bisulfate salts useful for the ΤI treatment of stress-related affective disorders

Gittos, Maurice Ward IN

PA Fr.

SO Brit. UK Pat. Appl., 26 pp.

CODEN: BAXXDU

DTPatent

English LA

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PAT	TENT	NO.					DATE			APPI	LICAT	ION I	NO.		D	ATE	
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		GM,	HR,	HU,	ID,	IL,	IN,	IS,	JΡ,	KE,	KG,	ΚP,	KR,	ΚZ,	LC,	LK,	LR,
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		ΝE,	SN,	TD,	TG												-
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ΕP	1420	788			A1		2004	0526		EP 2	2002-	7552	26		21	0020	822
ΕP	1420	788			В1		2006	1213									
	R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,
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BR	2002	0121	75		A		2004	0720		BR 2	2002-	1217	5	•	2	0020	822
CN	1549	715			Α		2004	1124		CN 2	2002-	8169	12		2	0020	822
JP	2005	5026	77		${f T}$		2005	0127		JP 2	2003-	5245	82		2	0020	822
							2005	0729		NZ 2	2002-	5313	45		2	0020	822
	PAT GB CA WO AU EP EP BR CN JP	GB 2379 CA 2459 WO 2003 W: RW: AU 2002 EP 1420 EP 1420 R: BR 2002 CN 1549 JP 2005	PATENT NO GB 2379216 CA 2459009 WO 20030202' W: AE, CO, GM, LS, PL, UA, RW: GH, CH, PT, NE, AU 20023215 EP 1420788 EP 1420788 EP 1420788 R: AT, IE, BR 20020121 CN 1549715	PATENT NO. GB 2379216 CA 2459009 WO 2003020275 W: AE, AG, CO, CR, GM, HR, LS, LT, PL, PT, UA, UG, RW: GH, GM, CH, CY, PT, SE, NE, SN, AU 2002321522 EP 1420788 EP 1420788 EP 1420788 EP 1420788 R: AT, BE, IE, SI, BR 2002012175 CN 1549715 JP 2005502677	PATENT NO. GB 2379216 CA 2459009 WO 2003020275 W: AE, AG, AL, CO, CR, CU, GM, HR, HU, LS, LT, LU, PL, PT, RO, UA, UG, US, RW: GH, GM, KE, CH, CY, CZ, PT, SE, SK, NE, SN, TD, AU 2002321522 EP 1420788 EP 1420788 EP 1420788 R: AT, BE, CH, IE, SI, LT, BR 2002012175 CN 1549715 JP 2005502677	PATENT NO. KIND DATE APPLICAT GB 2379216 A 20030305 GB 2001- CA 2459009 A1 20030313 WO 2002- W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, NE, SN, TD, TG AU 2002321522 A1 20030318 AU 2002- EP 1420788 B1 20040526 EP 2002- EP 1420788 B1 20061213 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BR 2002012175 A 20040720 BR 2002- CN 1549715 A 20041124 CN 2002- JP 2005502677 T 20050127 JP 2003-	PATENT NO.	DATE	PATENT NO.	PATENT NO. KIND DATE APPLICATION NO. DATE GB 2379216 A 20030305 GB 2001-20821 20030305	PATENT NO.						

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	US 7189742	B2	20070313		
	IN 2004MN00151	Α	20050624	IN 2004-MN151	20040227
PRAI	GB 2001-20821	Α	20010828		
	WO 2002-GB3869	W	20020822		
os	MARPAT 138:204953				
GT					

$$R^1$$
 $Me Me$
 $Me Me$
 $Me_2N(CH_2)_n$
 N
 $Me_2N(CH_2)_n$
 N

Ι

Title compds. [I; R1 = MeO, EtO, OH; R2 = H, R1; n = 2, 3], were prepared Thus, a cooled solution of H2SO4 in EtOH was mixed into 3-(3-methoxyphenyl)-3-(3-dimethylaminopropyl)-4,4-dimethylpiperidine-2,6-dione (AGN 2979) in EtOH followed by removal of solvent under reduced pressure and recrystn. from EtOH to give the bisulfate (II). II at 65 mg every 2 days in a 90 kg human male eliminated episodes of obstructive sleep apnea. II drug formulations are given. I are devoid of the weight loss and hepatocyte changes in the rat which limited to marginally effective levels the permitted clin. doses of the corresponding hydrochlorides in the treatment or prophylaxis of stress-related affective disorders such as anxiety, depression, migraine and sleep apnea.

IT 500350-77-6P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of piperidine-2,6-dione bisulfate salts useful for the treatment of stress-related affective disorders)

RN 500350-77-6 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl-, sulfate (1:1) (CA INDEX NAME)

CM 1

CRN 53873-21-5 CMF C19 H28 N2 O3

RE.CNT 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2007 ACS on STN

AN 1989:580731 CAPLUS

DN 111:180731

TI Anxiolytic pharmaceuticals containing 3-phenyl-3-(aminoalkyl)-4-methyl-2,6-dioxopiperidine derivatives

IN Costall, Brenda

PA National Research Development Corp., UK

SO Brit. UK Pat. Appl., 45 pp.

CODEN: BAXXDU

DT Patent

LA English

FAN.CNT 1

L'ATA'	CIVI				
	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
ΡI	GB 2206491	A	19890111	GB 1988-16214	19880707
	GB 2206491	В	19910123		
	EP 299680	A2	19890118	EP 1988-306208	19880707
	EP 299680	A3	19890726		
	R: AT, BE, CH,	DE, FR	GB, IT, LI	, LU, NL, SE	
	DK 8803826	Α	19890111	DK 1988-3826	19880708
	AU 8818862	A	19890112	AU 1988-18862	19880708
	AU 609496	B2	19910502		
	JP 01063517	A	19890309	JP 1988-173621	19880711
	ZA 8804986	A	19900328	ZA 1988-4986	19880711
PRAI	GB 1987-16338	Α	19870710		
OS GT	MARPAT 111:180731				

AB 3-Phenyl-3-aminoalkyl-4-methyl-2,6-dioxopiperidine derivs. (I; R1 = H, alkyl; n = 1, 2; R2 = H, Me, provided that one of R2 = H if n = 2; R3 = H, alkyl; R4 = alkyl; R5, R6 = H, Me; m = 0-3; each Y is in a meta or para position and represents OH, alkoxy, alkyl, hydroxyalkyl, halo, CF3, provided that OH and alkoxy are not in the para position) or their salts antagonize anxiogenesis associated with the withdrawal of addictive drugs, especially alc., nicotine, and cocaine. Tablets contained 3-(3'-methoxyphenyl)-

3-(3"-N,N-dimethylaminopropyl)-4,4-dimethyl-2,6-dioxopiperidine (II)

(base) 1, lactose 51.5, dried maize starch 45, and Mg stearate 1.5 mg/tablet. Mice were exposed to 8% alc. in the drinking water and during alc. withdrawal they received 10 mg diazepam/kg i.p. or 0.5 mg II/kg i.p. The mice were previously kept in a darkened box and during testing placed in a test area with white and black areas; during alc. intake the mice showed anxiolysis characterized by increased exploratory behavior in the white section and when the alc. was withdrawn the reverse profile was observed Both diazepam and II not only reversed anxiogenesis but actually led to anxiolysis; both appeared to be equieffective to combat anxiogenesis in alc. withdrawal, but II was more potent and devoid of the initial sedative action seen on treatment with diazepam. Both II and diazepam antagonized anxiogenesis in cocaine withdrawal in mice or in nicotine withdrawal in marmosets. I had no action on benzodiazepine receptors.

IT 53873-21-5 117576-37-1 123323-80-8

RL: BIOL (Biological study)

(as anxiolytic, for treatment of anxiogenesis associated with addictive drug withdrawal)

RN 53873-21-5 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl- (CA INDEX NAME)

RN 117576-37-1 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl-, (-)- (CA INDEX NAME)

Rotation (-).

RN 123323-80-8 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4dimethyl-, hydrochloride (9CI) (CA INDEX NAME)

Me (CH₂)
$$_3$$
 - NMe₂

●x HCl

EP 1988-306207

MARPAT 111:146823

os

GI

L13 ANSWER 4 OF 4 CAPLUS COPYRIGHT 2007 ACS on STN 1989:546823 CAPLUS AN111:146823 DN Phenyl-3-aminoalkyl-4-methyl-2,6-dioxopiperidine derivatives and their use TI as antipsychotic agents IN Costall, Brenda PΑ National Research Development Corp., UK SO Eur. Pat. Appl., 14 pp. CODEN: EPXXDW DTPatent LA English FAN.CNT 1 PATENT NO. KIND DATE APPLICATION NO. DATE -----_____ ----_____ _____ EP 1988-306207 19880707 PΙ EP 298738 A2 19890111 EP 298738 А3 19890809 EP 298738 B1 19920930 R: AT, BE, CH, DE, FR, GB, IT, LI, LU, NL, SE GB 2206490 Α 19890111 GB 1988-16213 19880707 GB 2206490 В 19910918 AT 81003 Т 19921015 AT 1988-306207 19880707 DK 8803825 19890111 DK 1988-3825 19880708 Α B1 19950814 DK 170360 AU 8818861 Α 19890127 AU 1988-18861 19880708 AU 606701 B2 19910214 ZA 8804937 Α 19900328 ZA 1988-4937 19880708 19890309 JP 1988-173620 19880711 JP 01063516 Α US 1988-217450 19880711 US 4877800 Α 19891031 CA 1988-571649 19880711 CA 1328077 С 19940329 PRAI GB 1987-16337 A. 19870710

19880707

Α

Ι

Phenyl-3-aminoalkyl-4-methyl-2,6-dioxopiperidine derivs. (I; R1, R3 = H, AB alkyl; n = 1,2; R2 = H, Me, provided that R2 = H when n = 2; R4 = alkyl; R5,R6 = H, Me; m = 0-3; Y is in a meta- or para-position; Y = OH, alkoxy, alkyl, hydroxyalkyl, halo, CF3, provided that OH and alkoxy are not in the para-position) or its salts are used for the manufacture of pharmaceuticals used in the treatment of psychosis. Hyperactivity was induced in rats via sterotaxic surgery, i.e. implantation of cannulae for intracerebral infusion of dopamine into the center of the nucleus accumbens and 25 µg dopamine was thus infused over a 24 h time period. Dopamine-induced hyperactivity occurred in a biphasic pattern between days 2-5 and 9-12 of treatment and could be antagonized with 0.01-10 mg/kg i.p. doses of 3-(3'-methoxyphenyl)-3-(3''-N, N-dimethylaminopropyl)-4,4-dimethyl-2,6dioxopiperidine (II); a lower dose of II (0.00001 mg/kg) controlled the 2nd peak but prevented control of the 1st peak. After withdrawal of II and dopamine a rebound of hyperactivity was not observed; persistent or excessive motor depression was not observed either with II during treatment. Fluphenazine at a 0.025-0.05 mg/kg dose was also effective in controlling dopamine-induced hyperactivity, however, after withdrawal, a rebound activity was observed Tablets contained II 1, lactose 51.5, dried maize starch 45, and Mg stearate 1.5 mg each.

IT 53873-21-5 117576-37-1 RL: BIOL (Biological study) (antipsychotic agent)

RN 53873-21-5 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl- (CA INDEX NAME)

RN 117576-37-1 CAPLUS
CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4dimethyl-, (-)- (CA INDEX NAME)

Rotation (-).

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666175-74-2 or 53873-21-5 or 92519-16-9 or
                                                      117576-37-1
             1 666175-74-2
                  (666175-74-2/RN)
             1 53873-21-5
                 (53873-21-5/RN)
             1 92519-16-9
                  (92519-16-9/RN)
             1 117576-37-1
                  (117576-37-1/RN)
L4
             4 666175-74-2 OR 53873-21-5 OR 92519-16-9 OR
                                                                117576-37-1
=> d 14
L4
     ANSWER 1 OF 4 REGISTRY COPYRIGHT 2007 ACS on STN
     666175-74-2 REGISTRY
RN
ED
     Entered STN: 22 Mar 2004
CN
     2,6-Piperidinedione, 3-(3,5-dimethoxyphenyl)-3-[3-(dimethylamino)propyl]-
     4,4-dimethyl-, (-)- (CA INDEX NAME)
FS
     STEREOSEARCH
MF
     C20 H30 N2 O4
CI
     COM
SR
     CA
LC
     STN Files:
                  CA, CAPLUS
Rotation (-).
          Me
            Me
                    NMe2
  HN
             (CH<sub>2</sub>) 3
 MeO
                OMe
**PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT**
               1 REFERENCES IN FILE CA (1907 TO DATE)
               1 REFERENCES IN FILE CAPLUS (1907 TO DATE)
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L4
     ANSWER 2 OF 4 REGISTRY COPYRIGHT 2007 ACS on STN
RN
     117576-37-1 REGISTRY
     Entered STN: 18 Nov 1988
ED
     2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-
CN
     dimethyl-, (-)- (CA INDEX NAME)
FS
     STEREOSEARCH
MF
     C19 H28 N2 O3
CI
     COM
SR
     CA
                  BEILSTEIN*, CA, CAPLUS, USPATFULL
     STN Files:
         (*File contains numerically searchable property data)
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Rotation (-).

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 6 REFERENCES IN FILE CA (1907 TO DATE)
- 6 REFERENCES IN FILE CAPLUS (1907 TO DATE)
- L4 ANSWER 3 OF 4 REGISTRY COPYRIGHT 2007 ACS on STN
- RN 92519-16-9 REGISTRY
- ED Entered STN: 17 Dec 1984
- CN 2,6-Piperidinedione, 3-(3,5-dimethoxyphenyl)-3-[3-(dimethylamino)propyl]-4,4-dimethyl- (CA INDEX NAME)
- MF C20 H30 N2 O4
- CI COM
- LC STN Files: CA, CAPLUS, USPAT7, USPATFULL

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PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 4 REFERENCES IN FILE CA (1907 TO DATE)
- 4 REFERENCES IN FILE CAPLUS (1907 TO DATE)
- L4 ANSWER 4 OF 4 REGISTRY COPYRIGHT 2007 ACS on STN
- RN 53873-21-5 REGISTRY
- ED Entered STN: 16 Nov 1984
- CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl- (CA INDEX NAME)

OTHER NAMES:

- CN 3-[3-(Dimethylamino)propyl]-3-(m-methoxyphenyl)-4,4-dimethyl-2,6-piperidinedione
- DR 117539-16-9
- MF C19 H28 N2 O3
- CI COM
- LC STN Files: BEILSTEIN*, CA, CAPLUS, IFICDB, IFIPAT, IFIUDB, MEDLINE,

PHAR, PROUSDDR, RTECS*, TOXCENTER, USPATFULL (*File contains numerically searchable property data)

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 11 REFERENCES IN FILE CA (1907 TO DATE)
- 11 REFERENCES IN FILE CAPLUS (1907 TO DATE)

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 L5
             12 L4
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         150877 WEIGHT
              0 L5 AND WEIGHT
 L6
 => d bib abs hitstr 15 1-12
      ANSWER 1 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN
 AN
      2007:329586 CAPLUS
 DN
      146:330838
      4,4-Dimethylpiperidine-2,6-dione derivatives for use in the treatment of
 ΤI
      hypertension
 IN
      Gittos, Maurice Ward
      Prestwick Pharmaceuticals, Inc., USA
 PA
 SO
      PCT Int. Appl., 20pp.
      CODEN: PIXXD2
 DT
      Patent
 LA
      English
 FAN.CNT 1
      PATENT NO.
                           KIND
                                  DATE
                                               APPLICATION NO.
                                                                       DATE
      ______
                           ____
                                  _____
                                               -----
 PΙ
      WO 2007031737
                            A1
                                  20070322
                                               WO 2006-GB3379
                                                                       20060913
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              CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
              GE, GH, GM, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP,
              KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN,
              MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS,
              RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ,
              UA, UG, US, UZ, VC, VN, ZA, ZM, ZW
          RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
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               GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
              KG, KZ, MD, RU, TJ, TM
 PRAI GB 2005-18763
                            Α
                                  20050914
 os
      MARPAT 146:330838
 AB
      Hypertension is treated with certain 3-phenyl-3-dimethylaminoalkyl-4,4-
      dimethylpiperidin-2,6-diones. The preferred compds. are
      3(3,5-dimethoxyphenyl)-3- (3-dimethylaminopropyl)-4,4-dimethylpiperidine-
      2,6-dione salts and, especially, 3(3-methoxyphenyl)-3-(3-dimethylaminopropyl)-
      4,4-dimethylpiperidine-2,6-dione salts.
 IT
      53873-21-5 92519-16-9 117576-37-1
      666175-74-2
      RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
       (Biological study); USES (Uses)
          (dimethylpiperidinedione derivs. for treatment of hypertension)
 RN
      53873-21-5 CAPLUS
 CN
      2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-
      dimethyl- (CA INDEX NAME)
         Me
```

RN 92519-16-9 CAPLUS

CN 2,6-Piperidinedione, 3-(3,5-dimethoxyphenyl)-3-[3-(dimethylamino)propyl]-4,4-dimethyl- (CA INDEX NAME)

RN 117576-37-1 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl-, (-)- (CA INDEX NAME)

Rotation (-).

RN 666175-74-2 CAPLUS

CN 2,6-Piperidinedione, 3-(3,5-dimethoxyphenyl)-3-[3-(dimethylamino)propyl]-4,4-dimethyl-, (-)- (CA INDEX NAME)

Rotation (-).

RE.CNT 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 2 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN AN 2003:174547 CAPLUS

```
DN
      138:204953
. TI
      Preparation of piperidine-2,6-dione bisulfate salts useful for the
      treatment of stress-related affective disorders
IN
      Gittos, Maurice Ward
PA
SO
      Brit. UK Pat. Appl., 26 pp.
      CODEN: BAXXDU
DT
      Patent
      English
LA
FAN.CNT 1
                           KIND
      PATENT NO.
                                  DATE
                                               APPLICATION NO.
                                                                       DATE
 PΙ
                            Α
                                  20030305
                                               GB 2001-20821
      GB 2379216
                                                                       20010828
      CA 2459009
                            Α1
                                  20030313
                                               CA 2002-2459009
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      WO 2003020275
                            A1
                                  20030313
                                               WO 2002-GB3869
                                                                       20020822
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              CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
              GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
              LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
              PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
              UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
          RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
              CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
              PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
              NE, SN, TD, TG
      AU 2002321522
                            A1
                                  20030318
                                               AU 2002-321522
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      EP 1420788
                            A1
                                  20040526
                                               EP 2002-755226
                                                                       20020822
                            В1
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      EP 1420788
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              IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK
      BR 2002012175
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                                  20040720
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      CN 1549715
                            Α
                                  20041124
                                               CN 2002-816912
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      JP 2005502677
                            \mathbf{T}
                                  20050127
                                               JP 2003-524582
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      NZ 531345
                            Α
                                  20050729
                                               NZ 2002-531345
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      AT 347891
                            Т
                                  20070115
                                               AT 2002-755226
                                                                       20020822
      ES 2275898
                            Т3
                                  20070616
                                               ES 2002-2755226
                                                                       20020822
      MX 2004PA01777
                            Α
                                  20041122
                                               MX 2004-PA1777
                                                                       20040225
      US 2004249159
                            A1
                                  20041209
                                               US 2004-486925
                                                                       20040225
                            В2
                                  20070313
      US 7189742
      IN 2004MN00151
                            Α
                                  20050624
                                               IN 2004-MN151
                                                                       20040227
 PRAI GB 2001-20821
                            Α
                                  20010828
```

20020822

WO 2002-GB3869

MARPAT 138:204953

os

GΙ

W

Ι

AB Title compds. [I; R1 = MeO, EtO, OH; R2 = H, R1; n = 2, 3], were prepared Thus, a cooled solution of H2SO4 in EtOH was mixed into 3-(3-methoxyphenyl)-3-(3-dimethylaminopropyl)-4,4-dimethylpiperidine-2,6-dione (AGN 2979) in EtOH followed by removal of solvent under reduced pressure and recrystn.

from EtOH to give the bisulfate (II). II at 65 mg every 2 days in a 90 kg human male eliminated episodes of obstructive sleep apnea. II drug formulations are given. I are devoid of the weight loss and hepatocyte changes in the rat which limited to marginally effective levels the permitted clin. doses of the corresponding hydrochlorides in the treatment or prophylaxis of stress-related affective disorders such as anxiety, depression, migraine and sleep apnea.

IT 92519-16-9

RL: RCT (Reactant); RACT (Reactant or reagent)
(preparation of piperidine-2,6-dione bisulfate salts useful for the treatment of stress-related affective disorders)

RN 92519-16-9 CAPLUS

CN 2,6-Piperidinedione, 3-(3,5-dimethoxyphenyl)-3-[3-(dimethylamino)propyl]-4,4-dimethyl- (CA INDEX NAME)

$$\begin{array}{c|c} & H & O \\ \hline & N & O \\ \hline & Me & & (CH_2)_3 - NMe_2 \\ \hline & Me & & OMe \\ \hline & MeO & & \end{array}$$

RE.CNT 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 3 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN

AN 1992:106095 CAPLUS

DN 116:106095

TI Process for preparation of 3-aryl-3-aminoalkyl-2,6-dioxohexahydropyridines

IN Dygos, John Henry; McLaughlin, Kathleen Therese; Ng, John Sau Hoi; Paul, Kalidas

PA G.D. Searle and Co., USA

SO Eur. Pat. Appl., 15 pp. CODEN: EPXXDW

DT Patent

LA English

FAN.CNT 1

1744.	ONT I				
	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
					
PΙ	EP 448972	A2	19911002	EP 1991-102833	19910226
	EP 448972	A3	19920506		
	R: AT, BE, CH,	DE, DK	, ES, FR,	GB, GR, IT, LI, LU, NL,	SE
	US 5104990	Α	19920414	US 1990-486027	19900227
	CA 2036968	A 1	19910828	CA 1991-2036968	19910225
	JP 04211657	Α	19920803	JP 1991-30929	19910226
	JP 06094460	В	19941124		
	US 5220019	Α	19930615	US 1992-859189	19920327
PRAI	US 1990-486027	Α	19900227		
os	CASREACT 116:106095	; MARPA	т 116:1060	95	
GI					•

AB A process is disclosed for the preparation of title compds. I [A = straight or branched C2-6 alkalene; R,R1 = C1-10 alkyl; Ar = heterocyclyl, (substituted) aryl] and particularly 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl-2,6-piperidinedione monohydrochloride (II), which is useful as an antidepressant. Thus, 6.58 Kg [(dimethylamino)propyl]methoxybenzeneacetonitrile, preparation given from 3-MeOC6H4CH2CN and C1(CH2)3NMe2.HCl, was treated with 11.89 Kg of (Me2CH)2CHLi and then 6.25 Kg Me2C:C(CO2Et)2 in THF-heptane to give 84.69% 3-MeOC6H4C(CN)[CMe2CH(CO2Et2](CH2)3NMe2.HCl. Hydrolysis of the latter compound in refluxing 96% H2SO4 and then treatment with 29% NH4OH followed by 36% aq HCl in EtOH gave 83.5% II.

IT 53873-21-5P

RL: SPN (Synthetic preparation); PREP (Preparation)
 (preparation of)

RN 53873-21-5 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl- (CA INDEX NAME)

L5 ANSWER 4 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN

AN 1991:609 CAPLUS

DN 114:609

TI Low-dosage anxiolytic compositions containing dioxopiperidine derivatives

IN Costall, Brenda

PA National Research Development Corp., UK

SO S. African, 56 pp.

CODEN: SFXXAB

DT Patent

LA English

FAN.CNT 1

1741. CIVI						
P.P	TENT NO.	KIND	DATE	APPLICATION NO.	DATE	
PI ZA	8804938	Α	19900328	ZA 1988-4938	19880708	
II	87059	Α	19921201	IL 1988-87059	19880710	
PRAI GE	3 1987-16340	Α	19870710		•	
os MA	RPAT 114:609					
CT						

Dioxopiperidine derivs. I [R1 = H, C1-4 alkyl; R2 = H, Me, provided that one R2 = H when n = 2; n = 1,2; R3 = H, C1-2 alkyl; R4 = C1-2 alkyl; R5, R6 = H, Me; m = 0-3; Y = OH, C1-2 alkoxy, C1-2 (hydroxy)alkyl, halo, trifluoromethyl in a meta or para position, provided that OH and alkoxy are not in the para position] or pharmaceutically acceptable salts are low-dosage anxiolytics; pharmaceutical compns. comprise I at 10-7-10-1 mg/unit dose. 3-(3'-Methoxyphenyl)-3-(3''-N,N-dimethylethylaminopropyl)-4,4-dimethyl-2,6-dioxopiperidine (II) at 0.00001-100.0 mg/kg s.c. showed anxiolytic activity in male albino BKW mice. The effect was achieved in the absence of sedation. Tablets comprise II 0.1, lactose 51.5, maize starch 45, and Mg stearate 1.5 mg/tablet.

IT 53873-21-5 117576-37-1

RL: BIOL (Biological study) (low-dosage anxiolytic)

RN 53873-21-5 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl- (CA INDEX NAME)

RN 117576-37-1 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl-, (-)- (CA INDEX NAME)

Rotation (-).

AN 1989:580731 CAPLUS

DN 111:180731

TΙ Anxiolytic pharmaceuticals containing 3-phenyl-3-(aminoalkyl)-4-methyl-2,6dioxopiperidine derivatives

IN Costall, Brenda

National Research Development Corp., UK PA

SO Brit. UK Pat. Appl., 45 pp. CODEN: BAXXDU

DTPatent

English LΑ

FAN.	CNT 1				
	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
ΡI	GB 2206491	Α	19890111	GB 1988-16214	19880707
	GB 2206491	В	19910123		
	EP 299680	A2	19890118	EP 1988-306208	19880707
	EP 299680	A3	19890726		
	R: AT, BE, CH,	DE, FR	GB, IT, L	I, LU, NL, SE	
	DK 8803826	Α	19890111	DK 1988-3826	19880708
	AU 8818862	Α	19890112	AU 1988-18862	19880708
	AU 609496	В2	19910502		
	JP 01063517	Α	19890309	JP 1988-173621	19880711
	ZA 8804986	Α	19900328	ZA 1988-4986	19880711
PRAI	GB 1987-16338	Α	19870710	·	•
os	MARPAT 111:180731				
GT				•	

$$R^{5}$$
 Me R^{6}
 R^{3} N (CH) n^{1} H2C N R^{1}

AB 3-Phenyl-3-aminoalkyl-4-methyl-2,6-dioxopiperidine derivs. (I; R1 = H, alkyl; n = 1, 2; R2 = H, Me, provided that one of R2 = H if n = 2; R3 = H, alkyl; R4 = alkyl; R5, R6 = H, Me; m = 0-3; each Y is in a meta or para position and represents OH, alkoxy, alkyl, hydroxyalkyl, halo, CF3, provided that OH and alkoxy are not in the para position) or their salts antagonize anxiogenesis associated with the withdrawal of addictive drugs, especially alc., nicotine, and cocaine. Tablets contained

3-(3'-methoxyphenyl)-3-(3"-N,N-dimethylaminopropyl)-4,4-dimethyl-2,6-dioxopiperidine (II) (base) 1, lactose 51.5, dried maize starch 45, and Mg stearate 1.5 mg/tablet. Mice were exposed to 8% alc. in the drinking water and during alc. withdrawal they received 10 mg diazepam/kg i.p. or 0.5 mg II/kg i.p. The mice were previously kept in a darkened box and during testing placed in a test area with white and black areas; during alc. intake the mice showed anxiolysis characterized by increased exploratory behavior in the white section and when the alc. was withdrawn the reverse profile was observed Both diazepam and II not only reversed anxiogenesis but actually led to anxiolysis; both appeared to be equieffective to combat anxiogenesis in alc. withdrawal, but II was more potent and devoid of the initial sedative action seen on treatment with diazepam. Both II and diazepam antagonized anxiogenesis in cocaine withdrawal in mice or in nicotine withdrawal in marmosets. I had no action on benzodiazepine receptors.

IT 53873-21-5 117576-37-1

RL: BIOL (Biological study)

(as anxiolytic, for treatment of anxiogenesis associated with addictive drug withdrawal)

RN 53873-21-5 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl- (CA INDEX NAME)

RN 117576-37-1 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl-, (-)- (CA INDEX NAME)

Rotation (-).

L5 ANSWER 6 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN

AN 1989:546823 CAPLUS

DN 111:146823

TI Phenyl-3-aminoalkyl-4-methyl-2,6-dioxopiperidine derivatives and their use as antipsychotic agents

IN Costall, Brenda

PA National Research Development Corp., UK

SO Eur. Pat. Appl., 14 pp.

CODEN: EPXXDW

DT Patent

LA English

FAN.CNT 1

	PAT	CENT	NO.			KINI)	DATE		A	PPLICAT	ON NO	o.	DATE
PΙ	ΕP	2987	38			A2		1989	0111	E	1988-	-30620	7	19880707
	EΡ	2987	38			A3		1989	0809					
	ΕP	2987	38			В1		1992	0930					
		R:	AT,	BE,	CH,	DE,	FR	, GB,	ΙΤ,	LI, I	U, NL,	SE		
	GB	2206	490			Α		1989	0111	GE	3 1988-	16213		19880707
	GB	2206	490			В		1991	0918					
	ΑT	8100	3			\mathbf{T}		1992	1015	A)	1988-	-30620	7	19880707
	DK	8803	825			Α		1989	0111	DF	1988-	3825		19880708

	DK 170360	В1	19950814		
•	AU 8818861	Α	19890127	AU 1988-18861	19880708
	AU 606701	B2	19910214		
	ZA 8804937	Α	19900328	ZA 1988-4937	19880708
	JP 01063516	Α	19890309	JP 1988-173620	19880711
	US 4877800	Α	19891031	US 1988-217450	19880711
	CA 1328077	С	19940329	CA 1988-571649	·19880711
PRAI	GB 1987-16337	Α	19870710		
	EP 1988-306207	Α	19880707		•
os	MARPAT 111:146823				
GI					

$$R^{5}$$
 Me R^{6}
 Y_{m}
 N_{m}
 N_{m}

AB Phenyl-3-aminoalkyl-4-methyl-2,6-dioxopiperidine derivs. (I; R1, R3 = H, alkyl; n = 1,2; R2 = H, Me, provided that R2 = H when n = 2; R4 = alkyl; R5, R6 = H, Me; m = 0-3; Y is in a meta- or para-position; Y = OH, alkoxy, alkyl, hydroxyalkyl, halo, CF3, provided that OH and alkoxy are not in the para-position) or its salts are used for the manufacture of pharmaceuticals used in the treatment of psychosis. Hyperactivity was induced in rats via sterotaxic surgery, i.e. implantation of cannulae for intracerebral infusion of dopamine into the center of the nucleus accumbens and 25 µg dopamine was thus infused over a 24 h time period. Dopamine-induced hyperactivity occurred in a biphasic pattern between days 2-5 and 9-12 of treatment and could be antagonized with 0.01-10 mg/kg i.p. doses of 3-(3'-methoxyphenyl)-3-(3''-N,N-dimethylaminopropyl)-4,4-dimethyl-2,6dioxopiperidine (II); a lower dose of II (0.00001 mg/kg) controlled the 2nd peak but prevented control of the 1st peak. After withdrawal of II and dopamine a rebound of hyperactivity was not observed; persistent or excessive motor depression was not observed either with II during treatment. Fluphenazine at a 0.025-0.05 mg/kg dose was also effective in controlling dopamine-induced hyperactivity, however, after withdrawal, a rebound activity was observed Tablets contained II 1, lactose 51.5, dried maize starch 45, and Mg stearate 1.5 mg each.

IT 53873-21-5 117576-37-1

RL: BIOL (Biological study) (antipsychotic agent)

Ι

RN 53873-21-5 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4dimethyl- (CA INDEX NAME)

RN 117576-37-1 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4dimethyl-, (-)- (CA INDEX NAME)

Rotation (-).

L5ANSWER 7 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN

1989:540496 CAPLUS ΑN

111:140496 DN

2,6-Piperidinediones as analgesics ΤI

IN Roberts, Malcolm Henry Traffod

PΑ National Research Development Corp., UK

SO Eur. Pat. Appl., 22 pp.

CODEN: EPXXDW

DT Patent

English LΑ

FAN.	CNT	1														
	PAT	TENT NO	•			KINI) -	DATE			APE	PLICATI	ON N	ю.]	DATE
PI	EP	295836				A2		1988	1221		EP	1988-3	0531	.7	- :	19880610
	EP	295836				A3		1989	0719							
	ΕP	295836				В1		1992	0902							
		R: A'	Γ,]	ΒE,	CH,	DE,	FR,	GB,	IT,	LI,	LU	J, NL,	SE			
	GB	2205745	5			Α		1988	1221		GB	1988-1	3796	;		19880610
	GB	2205745	5			В		1990	0919							
	ΑT	80035				T		1992	0915		AΤ	1988-3	0531	.7 .		19880610
	AU	8817676	6			Α		1988	1222		AU	1988-1	7676	j	:	19880614
	AU	606424				B2		1991	0207							
	US	4871750	0			Α		1989	1003		US	1988-2	0627	3		19880614
	DK	8803282	2			Α		1988	1217		DK	1988-3	282			19880615
	ZA	8804275	5			Α		1989	0530		ZA	1988-4	275			19880615
	JP	0101676	63			Α		1989	0120		JΡ	1988-1	4923	7	:	19880616
PRAI	GB	1987-14	403	3		Α		1987	0616							
	GB	1987-14	437	4		Α		1987	0619							
	EΡ	1988-30	053	17		Α		1988	0610			ı				
OS GI	MAI	RPAT 11	1:1	4049	96											

Phenyl-3-(aminoalkyl)-4-methyl-2,6-piperidinediones I (R1 = H, C1-4 alkyl; R2 = H, Me with one R2 = H when n = 2; R3 = H, Me, Et; R4 = Me, Et; R5, R6 = H, Me; Y = OH, MeO, EtO, Me, Et, HOCH2, hydroxyethyl, halo, CF3; n = 1, 2; m = 0-3 with each Y in a meta or para position) or their salts are useful as analgesics. Using the tail-flick latency test, (-)-3-(3-methoxyphenyl)-3-[3-(dimethylamino)propyl]-4,4-dimethyl-2,6-piperidinedione [(-)-II] injected into rats at 2 mg/kg had a strong analgesic effect with the peak response delayed until 20 min after the injection and baseline latencies were not recovered until 2 h after the injection; the potency was of the same order of magnitude as morphine with a similar time course of effect. Naloxone, known to block drug actions at opioid receptors, failed to reduce the potency of this compound Tablets contained II 50, lactose 51.5, dried corn starch 45, and Mg stearate 1.5 mg/tablet.

IT 53873-21-5 117576-37-1
RL: BIOL (Biological study)
(analgesic pharmaceuticals containing)

Ι

RN 53873-21-5 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl- (CA INDEX NAME)

RN 117576-37-1 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl-, (-)- (CA INDEX NAME)

Rotation (-).

L5 ANSWER 8 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN

AN 1988:622491 CAPLUS

DN 109:222491

TI Anxiolytic compositions containing dioxopiperidine derivatives

IN Gittos, Maurice Ward; Costall, Brenda

PA National Research Development Corp., UK

SO Eur. Pat. Appl., 34 pp. CODEN: EPXXDW

DT Patent

LA English

FAN.CNT 2

	PATENT NO.		DATE	APPLICATION NO.	DATE
PI	EP 263594 EP 263594 EP 263594	A3	19880413 19890802 19920624	EP 1987-307860	19870904
	=			LI, LU, NL, SE	
	GB 2181346	A B	19870423 19891004	GB 1986-21577	19860908
	GB 2196251 GB 2196251	A B	19880427 19900704	GB 1987-20813	19870904
	CA 1316112	C	19930413		19870904
	DK 8704654	Α	19880309		19870907
	AU 8778109	Α	19880310	AU 1987-78109 ,	19870907
	AU 602716	B2	19901025		
	JP 63101361	Α	19880506	JP 1987-225124	19870908
	AU 9059784	Α	19901101	AU 1990-59784	19900724
PRAI	GB 1986-21577	Α	19860908		
	GB 1987-16339	Α	19870710		
	GB 1985-22455	Α	19850911		
	GB 1986-3909	Α	19860217		
	GB 1986-3910	Α	19860217		
	GB 1987-16359	Α	19870710		•
OS GI	MARPAT 109:222491				

A pharmaceutical composition in unit dose form comprises, with a . AB pharmaceutically acceptable diluent or carrier, 10-7-10-1 mg/unit dose of 3-phenyl-3-aminoalkyl-4-methyl-2,6-dioxopiperidines I (R1 = H, C1-4 alkyl; n = 1,2; R2 = H, Me, provided that one R2 = H when n = 2; R3 = H, C1-2alkyl; R4 = C1-2 alkyl; R5, R6 = H, Me; m = 0-3; Y = OH, C1-2 alkoxy, C1-2alkyl, C1-2 hydroxyalkyl, halo, CF3, in meta or para position, provided that OH and alkoxy are not in para position) or pharmaceutically acceptable salts for treatment of anxiety. Native male albino BKW mice in an anti-anxiety test were administered 3(3'-methoxyphenyl)-3(3''-N,Ndimethylaminopropyl)-4,4-dimethyl-2,6-dioxopiperidine (II) in water by s.c. injection or diazepam in PEG and water by i.p. injection. II was as effective as diazepam and, in fact, was exceptionally potent (0.00001-100.0 mg/kg) and showed a dose range of 10 million (106). dose related effects of II contrasted with the all-or-none response of diazepam. A gelatin capsule formulation comprised II HCl 2.5 and talc 70 mg/capsule.

IT 53873-21-5 92519-16-9 117576-37-1 RL: BIOL (Biological study)

(anxiolytic)

RN 53873-21-5 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl- (CA INDEX NAME)

RN 92519-16-9 CAPLUS

CN 2,6-Piperidinedione, 3-(3,5-dimethoxyphenyl)-3-[3-(dimethylamino)propyl]-4,4-dimethyl- (CA INDEX NAME)

RN 117576-37-1 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl-, (-)- (CA INDEX NAME)

Rotation (-).

L5 ANSWER 9 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN

AN 1987:605181 CAPLUS

DN 107:205181

TI Use of dioxopiperidine derivatives in the treatment of anxiety, for the reduction of abnormally high brain levels of serotonin or 5-hydroxyindoleacetic acid, and in the treatment of bacterial or viral infections

IN Gittos, Maurice Ward

PA National Research Development Corp., UK

SO Eur. Pat. Appl., 38 pp.

CODEN: EPXXDW

DT Patent

LA English

FAN.CNT 2

GI

	PAT	TENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	EP	216555 216555 216555	A3	19891123	EP 1986-306920	19860908
		R: BE, CH, DE	FR, GE	, IT, LI,	LU, NL, SE	
					EP 1991-105508	19860908
	ΕP	452765				
		R: BE, CH, DE	•		•	
		8604337		19870312	DK 1986-4337	
		62061919			JP 1986-213704	19860910
		4738973			US 1986-905525	
		8662601			AU 1986-62601	19860911
	ΑU	588365	В2	19890914		
	CA	1273879	A1	19900911	CA 1986-518034	19860911
	US	4835151	Α	19890530	US 1987-136996	19871223
	US	4918084	Α	19900417	US 1989-323308	19890314
	US	4994475	Α	19910219	US 1989-452343	19891219
PRAI	GB	1985-22455	Α	19850911		
	GB	1986-3909	Α	19860217		
	GB	1986-3910	Α	19860217		
	US	1986-905525	A 3	19860910		
	US	1987-136996	A 3	19871223		
	US	1989-323308	A3	19890314		
os	MAI	RPAT 107:205181			•	

$$R^{1}$$
 R^{4}
 R^{5}
 NH
 R^{2}
 $(CH_{2})_{n}NR_{2}^{3}$

AB The title compds. I (R1 = OMe, OEt, OH; R2 = H, OMe, OEt, OH; R3 = Me, Et; R4, R5 = H, Me; n = 2, 3) or their pharmacol. acceptable acid addition salts are used in medications for treatment of anxiety or to counter the anxiogenic activity of benzodiazepine inverse agonists. They are also used for reduction of chronic high brain levels of serotonin or 5-hydroxyindoleacetic acid, or treatment of bacterial or viral infections. A tablet contained from I (R1 = OMe, R2 = R5 = H, R3 = R4 = Me, n = 3) (II) 100, Tranxene 10, wheat starch 7, lactose 20, and Mg stearate 1 mg. The anxiolytic activity of II in rats was between the activity of chlorodiazepoxide and diazepam, and its sedative effect was less than that of the benzodiazepines. In clin. tests the combination of II and Tranxene decreased anxiety in hospitalized depressive patients. At 120 mg/day, II stopped sleep apnea in a male patient.

IT 53873-21-5

RL: BIOL (Biological study)

Ι

(pharmaceutical, for treatment of anxiety)

RN 53873-21-5 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl- (CA INDEX NAME)

L5 ANSWER 10 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN

AN 1984:577535 CAPLUS

DN 101:177535

TI Treatment of migraine with dioxopiperidine derivatives

IN Gittos, Maurice W.; Amey, David A.

PA USA

SO U.S., 5 pp.

CODEN: USXXAM

DT Patent

LA English

FAN.CNT 1

PATE	ENT NO.	KIND	DATE	APPLICATION NO.	DATE
	1461771 1983-471099	Α	19840724 19830301	US 1983-471099	19830301

$$R^{1}$$
 R^{2}
 $R^{3}N(CH_{2})$
 $R^{3}N(CH_{2})$
 $R^{3}N(CH_{2})$

AB Migraine is treated or prevented with I derivs. (R1 and R2 = H, MeO, EtO, or HO, R3 = Me or Et and n = 2 or 3) or their salts. I(R1 = MeO, R2 = H, R3 = Me, n = 3).HCl (II) [53873-28-2] was prepared by intramol.condensation of Et 4-(3-N,N-dimethylaminopropyl)-4-cyano-4-(3-methoxyphenyl)-3,3-dimethylbutanoate <math>[53873-27-1] by refluxing in 2.5N HCl. Tablets were prepared containing 50 mg II each.

IT 53873-21-5P 92519-16-9P RL: PREP (Preparation)

Ι

(preparation of, for migraine headache pharmaceuticals)

RN 53873-21-5 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl- (CA INDEX NAME)

RN 92519-16-9 CAPLUS

CN 2,6-Piperidinedione, 3-(3,5-dimethoxyphenyl)-3-[3-(dimethylamino)propyl]-4,4-dimethyl- (CA INDEX NAME)

Me
$$(CH_2)_3 - NMe_2$$

Me OMe

L5 ANSWER 11 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN

AN 1975:564204 CAPLUS

DN 83:164204

OREF 83:25774h,25775a

TI Alkyl esters, dialkyl amides, and saturated heterocyclic amides of 4-aminoalkyl-4-cyano-4-phenylbutanoic and -but-2-enoic acids

IN Gittos, Maurice W.; Amey, David A.

PA Aspro-Nicholas Ltd., UK

SO Ger. Offen., 36 pp.

CODEN: GWXXBX

DT Patent

LA German

FAN.CNT 2

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	DE 2459077	A1	19750703	DE 1974-2459077	19741213
	GB 1458537	А	19761215	GB 1973-58202	19741203
	AU 7476155	А	19760610	AU 1974-76155	19741206
	ZA 7407769	Α	19760825	ZA 1974-7769	19741206
	US 3998965	Α	19761221	US 1974-531556	19741211
	BE 823272	A1	19750612	BE 1974-151436	19741212
	NL 7416161	Α	19750617	NL 1974-16161	19741212
	DK 7406522	Α	19750825	DK 1974-6522	19741213
	FR 2254329	A1	19750711	FR 1974-41315	19741216
	JP 50089343	A	19750717	JP 1974-144360	19741216
	US 4035497	Α	19770712	US 1976-679165	19760422
PRAI	GB 1973-58202	Α	19731215		
	GB 1972-59761	Α	19721228		
	US 1973-425876	A2	19731218		
	US 1974-531556	A3	19741211	,	

GI For diagram(s), see printed CA Issue.

Butenoate I, isolated as the H oxalate, was prepared by treating NaH and m-MeOC6H4CH(CN)CH2CH2NMe2 in Me2SO with PhSO2OCEt:C(CO2Et)2. Butanoic acid derivative II (R = R1 = R2 = Me, R3 = H, R4 = OEt, n = 3) (III) was prepared from m-MeOC6H4CH(CN)(CH2)3NMe2 and 3,3-dimethyl-1-ethoxyprop-2-enylidenemorpholinium tetrafluoroborate. II (R = PhCH2, Me; R1, R2 = H or Me; R3 = Me or H; R4 = morpholino, n = 2 or 3) were prepared from IV and the appropriate morpholinium tetrafluoroborate. II (R = R1 = R3 = Me, R2 = H, R4 = morpholino, n = 2) was prepared by treating Me2NCH2CH2C(CN)(C6H4OMe-m)CHMeCMe:C(OEt)R5 (R5 = morpholino) with MeSO3H and NaI in EtOH. I and III were cyclized to hydropyridines with NH3. I and II have antidepressant and cardiovascular activity (no data).

IT 53873-21-5P

RN 53873-21-5 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl- (CA INDEX NAME)

L5 ANSWER 12 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN

AN 1974:520488 CAPLUS

DN 81:120488

OREF 81:19043a,19046a

TI Antidepressant 3-(aminoalkyl)-3-phenyl-2,6-dioxopiperidines or -tetrahydropyridines

IN Gittos, Maurice W.; Amey, David A.

PA Aspro-Nicholas Ltd.

SO Ger. Offen., 52 pp.

CODEN: GWXXBX

DT Patent LA German

FAN.CNT 2

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI .	DE 2363052	A1	19740711	DE 1973-2363052	19731219
	DE 2363052	C2	19880721		
	AU 7363761	Α	19750619	AU 1973-63761	19731218
	US 3963729	Α	19760615	US 1973-425876	19731218
	ZA 7309598	Α	19741127	ZA 1973-9598	19731220
	BE 808958	A1	19740621	BE 1973-139144	19731221
	GB 1455687	Α	19761117	GB 1972-59761	19731227
	FR 2212147	A 1	19740726	FR 1973-46908	19731228
	JP 49094683	Α	19740909	JP 1974-4486	19731228
	JP 60053014	В	19851122		
	US 4035497	Α	19770712	US 1976-679165	19760422
PRAI	GB 1972-59761	Α	19721228		
	GB 1973-58202	Α	19731215		
	US 1973-425876	A2	19731218		
	US 1974-531556	A3	19741211		
~ =	- 1: / \		1 ~~ -		

GI For diagram(s), see printed CA Issue.

About 20 hydrogenated pyridines I (n = 2 or 3; R = Me2N, Et2N, or PhCH2NMe; R1 = H, 3-MeO, or 4-Cl; R2 = H, Me, or Et; R3 = H or Me; R4 = H, Me, or CO2Et) and II (R5 = Me or Et; R6 = H or CO2Et) or their salts were prepared I had antidepressant and minor parasympatholytic activity when tested i.p. in the rat. Thus, 3-MeOC6H4CH(CN)(CH2)nNMe2 (III, n = 3) was treated with NaH in Me2SO and with 4-(1-ethoxy-3,3-dimethyl-2-propenylidene)morpholinium tetrafluoroborate to give 3-MeOC6H4C(CN)[(CH2)3NMe2]CMe2CH:CR7 - OEt (R7 = morpholino), which was cyclized in H2SO4 and AcOH at 100° to give I (n = 3, R = Me2N, R1 = 3-MeO, R2 = R3 = Me, R4 = H). III (n = 2) was treated with NaH in Me2SO and with PhSO3CEt:C(CO2Et)2 to give 3-MeOC6H4C(CN) (CH2CH2NMe2)CEt:C(CO2Et)2, which on treatment with H2SO4 and AcOH at 100° gave II (R5 = Et, R6 = CO2Et).

IT 53873-21-5P

RL: SPN (Synthetic preparation); PREP (Preparation)
 (preparation of)

RN 53873-21-5 CAPLUS

CN 2,6-Piperidinedione, 3-[3-(dimethylamino)propyl]-3-(3-methoxyphenyl)-4,4-dimethyl- (CA INDEX NAME)